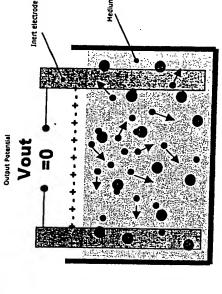
Title: I ransient Electrical Signal Based Methods and Devices for Characterizing Molecular Interaction and/or Motion in a Sample

and/or!Motion in a Sample Inventor: Pourmand, Nader Application No.: 10/040,030 Atty. Docket No. STAN-241

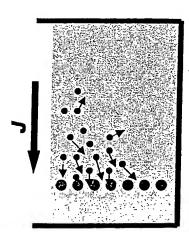
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J: Current Density of B

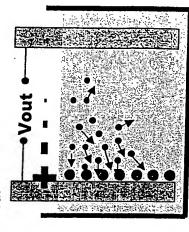
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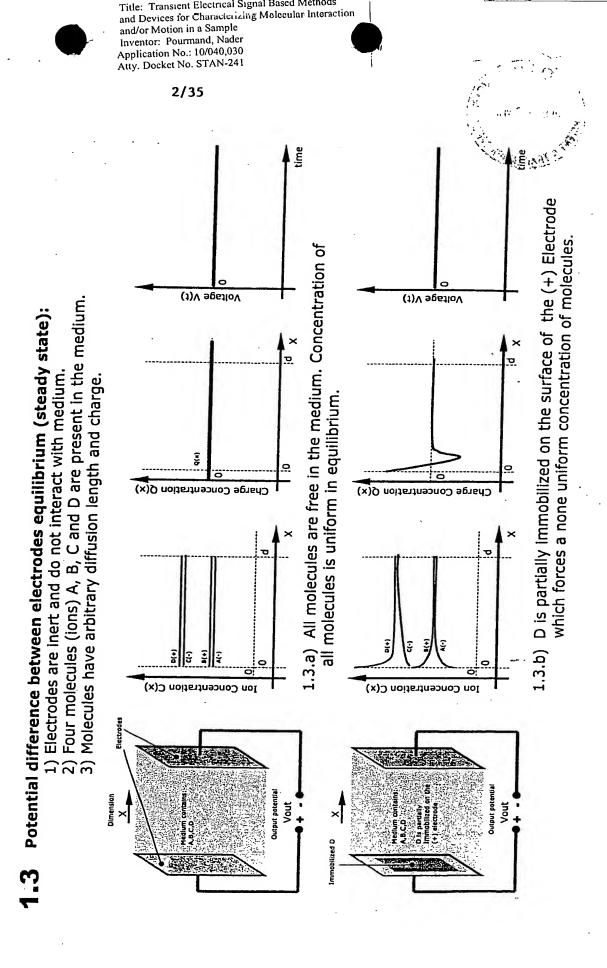


1.1) If A and B are both free in the medium no net transient gradient of molecules (current density) is created.



a net transient gradient (current density) of B toward A. This transient current 1.2) If A is spatially immobilized and B is free in the medium, the reaction causes creates a temporary potential difference in the medium.





Inventor: Pourmand, Nader Application No.: 10/040,030 Atty. Docket No. STAN-241 3/35 Potential difference between electrodes in none-equilibrium (transient) state: 1.4.b) C and D are added asymmetric to the electrodes and an ionic perturbation is generated. Two molecules (ions) are present in the medium and two are added. 1.4.a) C and D are added symmetric to the electrodes. (1)V agetioV Voltage V(t) Electrodes are inert and do not interact with the medium. Molecules have arbitrary diffusion length and charge. 4) Background molecules are not shown (A and B) Charge Concentration Q(x) Charge Concentration Q(x) T 7 t 0 € Ion Concentration C(x) Ion Concentration C(x) Output potential Vout Vout

Transient Electrical Signal Based Methods and

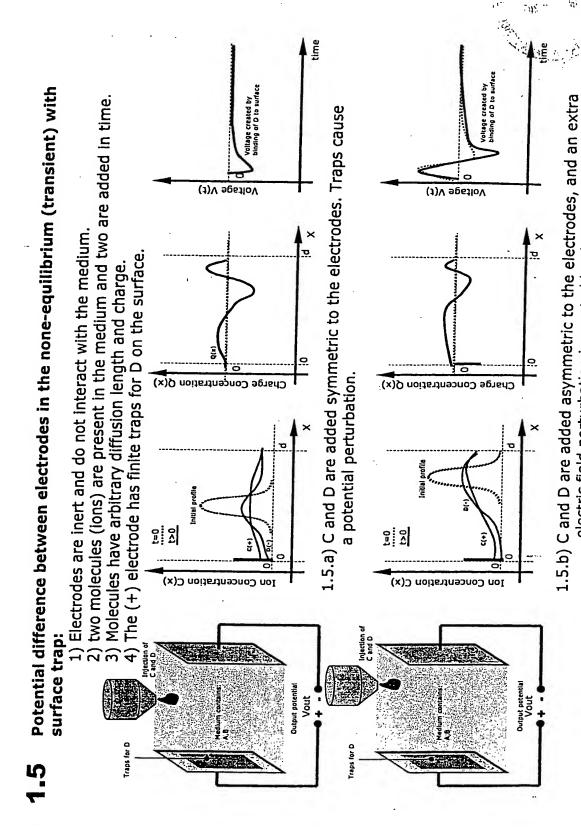
Devices for Characterizing Molecular Interaction

and/or Motion in a Sample

Title: Transient Electrical Signal Based Methods and Devices for Characterizing Molecular Interaction and/or Motion in a Sample

and/or Motion in a Sample Inventor: Pourmand, Nader Application No.: 10/040,030 Atty. Docket No. STAN-241

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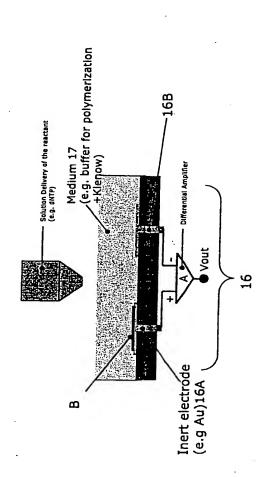


electric field perturbation is created by the traps.

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2.1 Planar sensor design example:

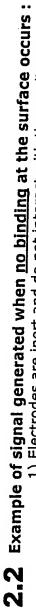
- 1) Electrodes are inert and do not interact with the medium.
 - 2) The target molecules are immobilized on the (+) electrode.
 - 3) The (-) electrode is the reference electrode.
- 4) A differential amplifier subtracts the voltage from the two electrodes.



and Devices for Characterizing Molecular Interaction and/or Motion in a Sample Inventor: Pourmand, Nader Application No.: 10/040,030 Atty. Docket No. STAN-241

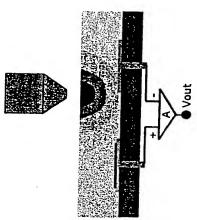
6/35

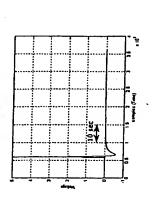




Electrodes are inert and do not interact with the medium.
The target molecules are immobilized on the (+) electrode.
The (-) electrode is the reference electrode.
A differential amplifier subtracts the voltage of the two electrodes.







a) Solution is delivered symmetric to the electrodes With 0.1 pmol immobilized ssDNA.

b) Solution is delivered asymmetric to the electrodes With 0.1 pmol immobilized ssDNA.

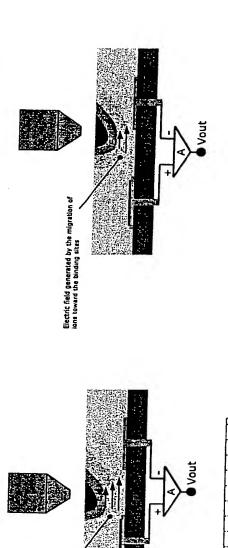
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2.3 Example of signal generated when binding at the surface occurs:

Electrodes are inert and do not interact with the medium.

The target molecules are immobilized on the (+) electrode. Electrodes are inert and do not interact with
The target molecules are immobilized on the
The (-) electrode is the reference electrode.

4) A differential amplifier subtracts the voltage from the two electrodes.

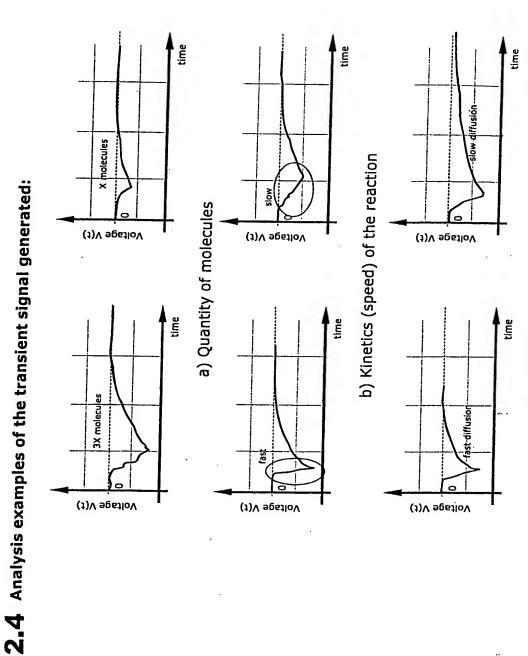


Electric field generated by the migration of ions toward the binding sites

b) Solution is delivered asymmetric to the electrodes Polymerization of 0.1 pmol primed ssDNA.

a) Solution is delivered symmetric to the electrodes, Polymerization of 0.1 pmol primed sSDNA.





a) Movement and diffusion of molecules.

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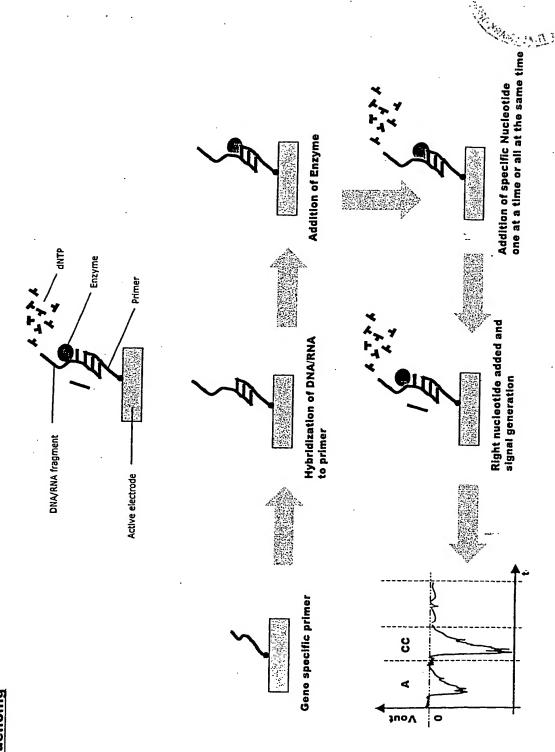
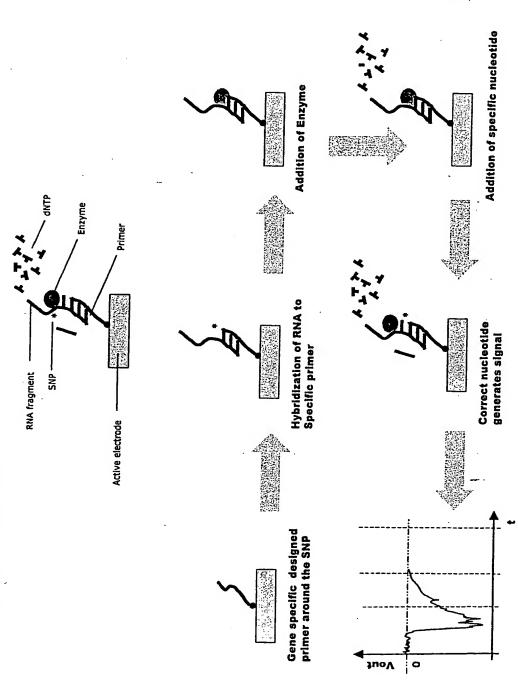


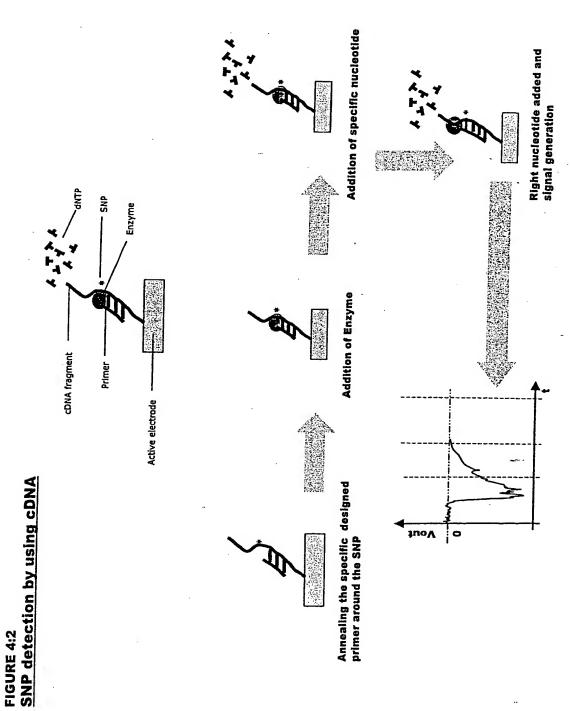
FIGURE 3: Sequencing and Devices for Characterizing Molecular Interaction and/or Motion in a Sample Inventor: Pourmand, Nader Application No.: 10/040,030 Atty. Docket No. STAN-241

FIGURE 4:1 SNP detection by using Total RNA



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SNP detection by using allele specific primer

FIGURE 4:3

Title: Transient Electrical Signal Based Methods and Devices for Characterizing Molecular Interaction and/or Motion in a Sample Inventor: Pourmand, Nader Application No.: 10/040,030 Atty. Docket No. STAN-241 12/35 Enzyme Addition of Enzyme & dNTP Hybridization of DNA/RNA to primer Perfect match DNA/RNA fragment miss-match Allele specific primers Active electrode Primer

Perfect match primer/template polymerizes and signal is generated

FIGURE 5: SNP detection by using PCR product Title: Transient Electrical Signal Based Methods and Devices for Characterizing Molecular Interaction and/or Motion in a Sample Inventor: Pourmand, Nader Application No.: 10/040,030 Atty. Docket No. STAN-241

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Addition of specific nucleotide Right nucleotide added and signal generation SNP. Enzyme Addition of Enzyme PCR fragment Primer Active electrode Annealing the specific designed primer around the SNP

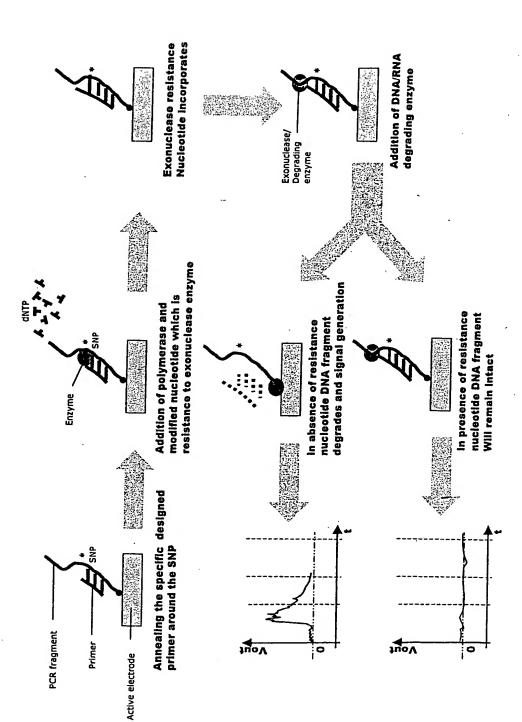
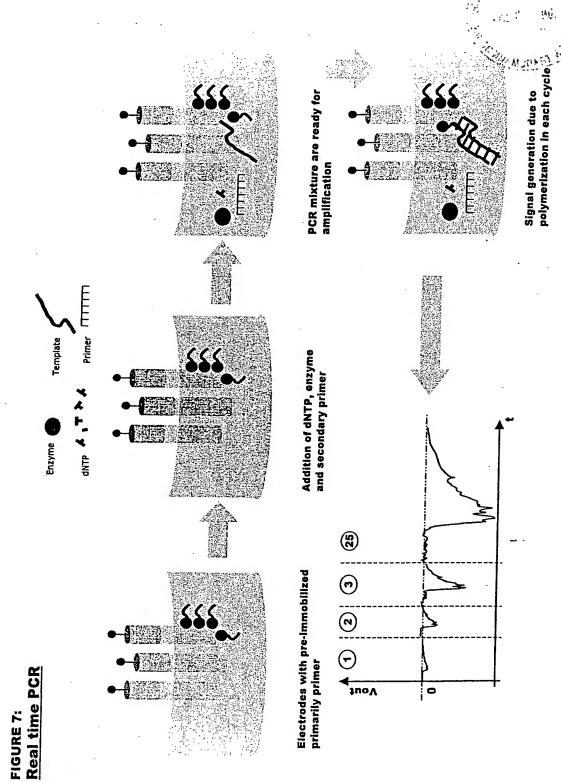


FIGURE 6: SNP detection by using Exonuclease/ Degrading enzyme

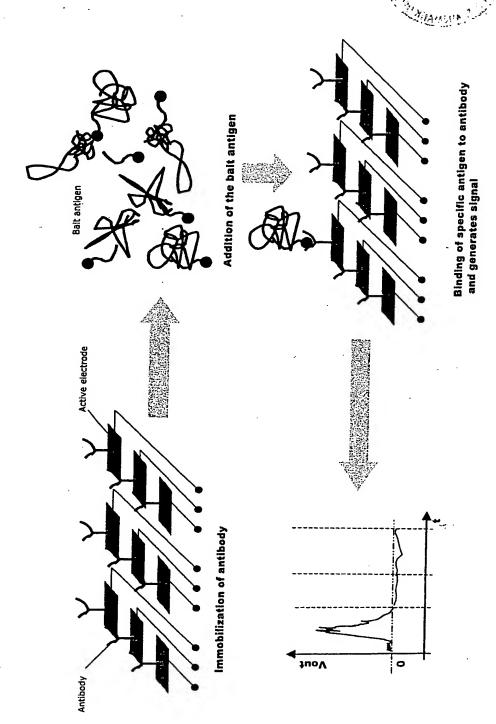
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Title: Transient Electrical Signal Based Methods and Devices for Characterizing Molecular Interact and/or Motion in a Sample Inventor: Pourmand, Nader Application No.: 10/040,030 Atty. Docket No. STAN-241 16/35 Addition of enzyme and dNTP Only the hybridized primer will polymerize and generates signal Denaturing and hybridization of pathogens DNA/RNA Pathogens DNA Active electrode 0 Immobilization of Pathogen specific primer

FIGURE 8: Pathogen typing





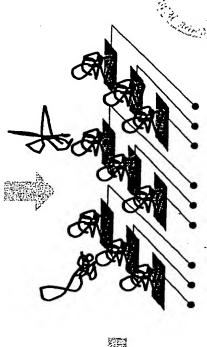
Antigen-antibody detection FIGURE 9:

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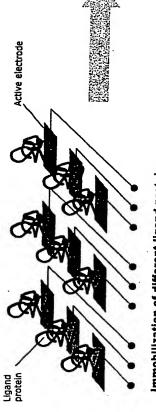




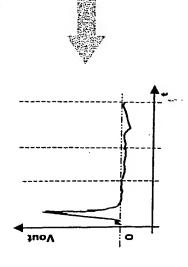
Addition of the bait proteins



Specific protein-protein interaction and generates signal







Protein-protein interaction FIGURE 10:

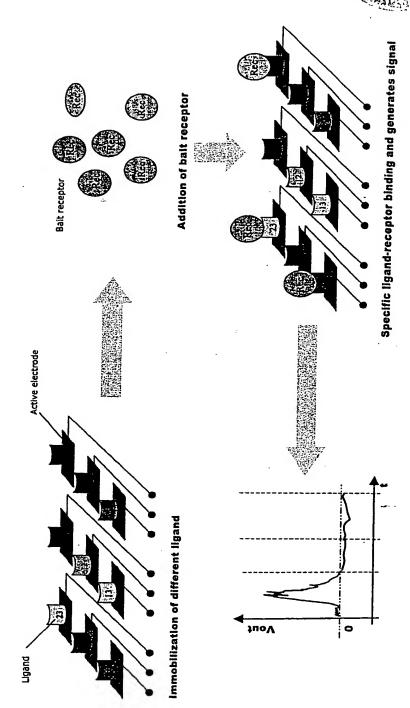


FIGURE 11: Ligand and receptor detection

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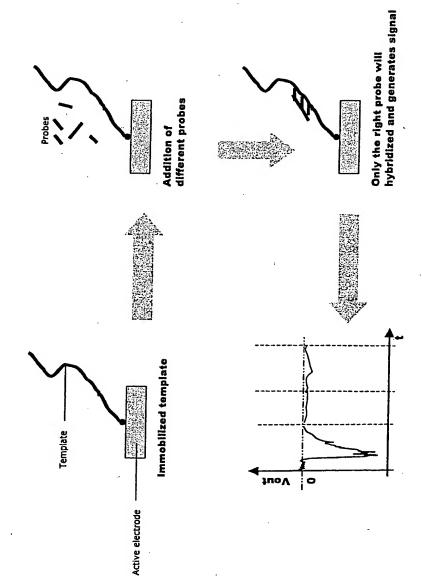
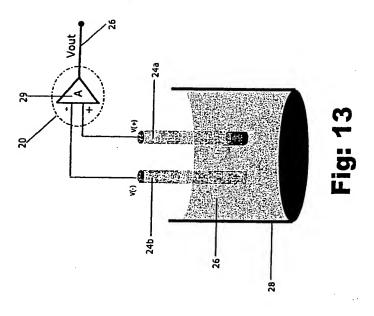
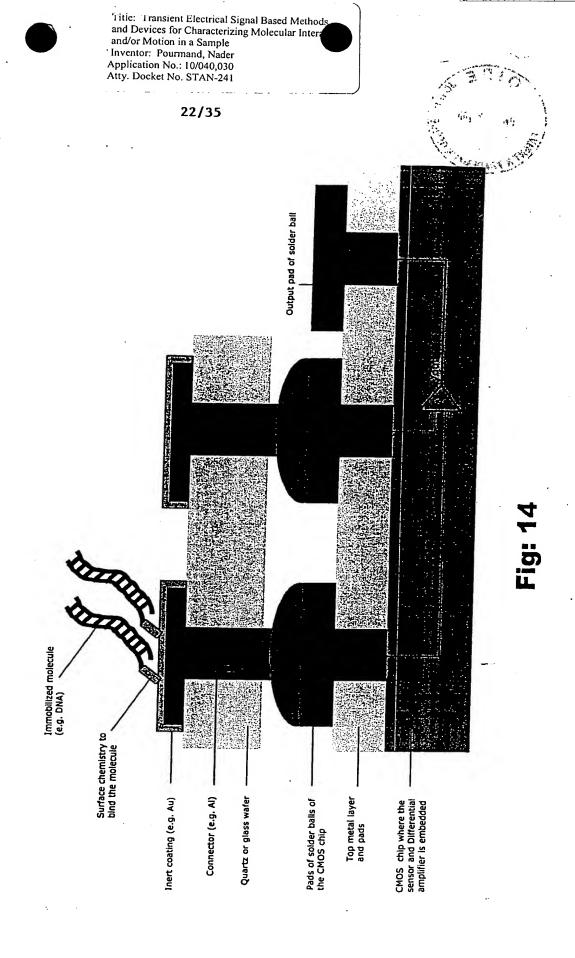


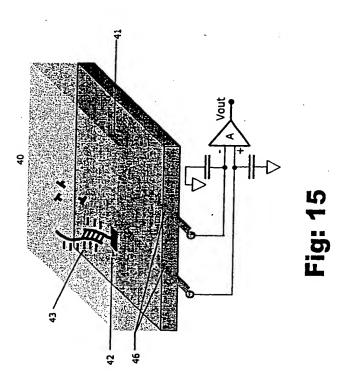
FIGURE 12: Hybridization



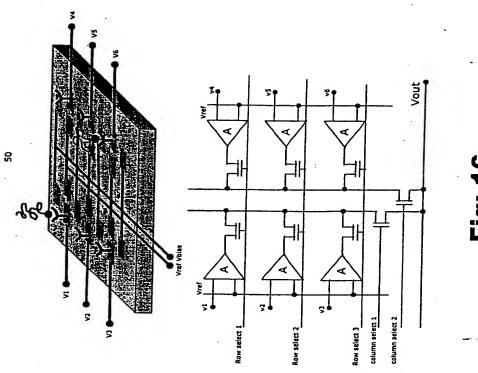






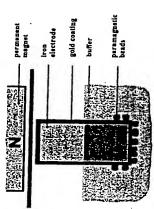






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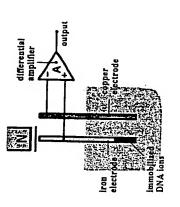


Figure 17 B: Basic model of the sensor with a differential amplifier



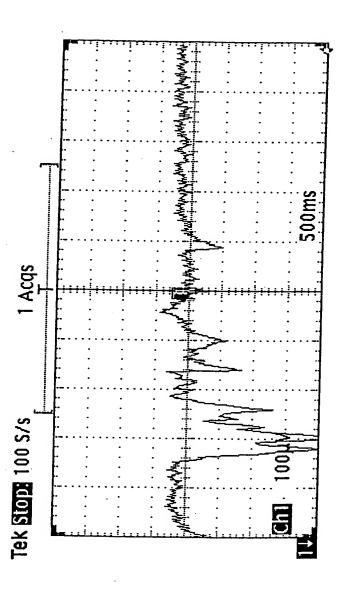
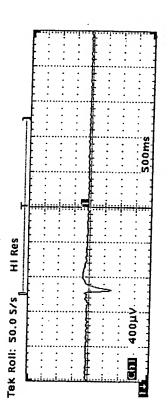


Figure 18 A: some sample charge sequencing extension signatures for 300 bp DNA





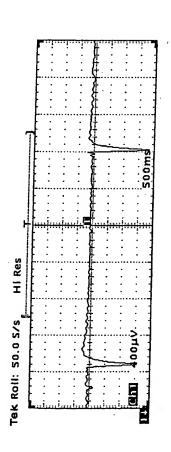
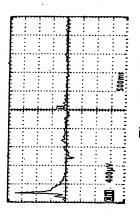


Figure 18 B: More sample charge sequencing extension signatures for 300 bp DNA with two different concentration of Immobilized DNA (0.05 pmol and 0.1 pmol)







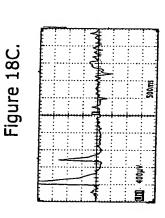
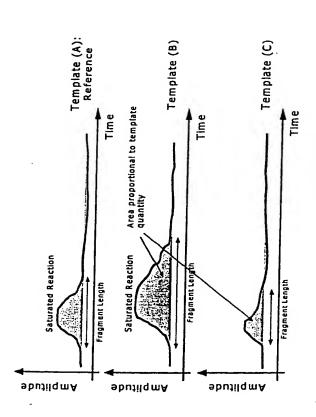
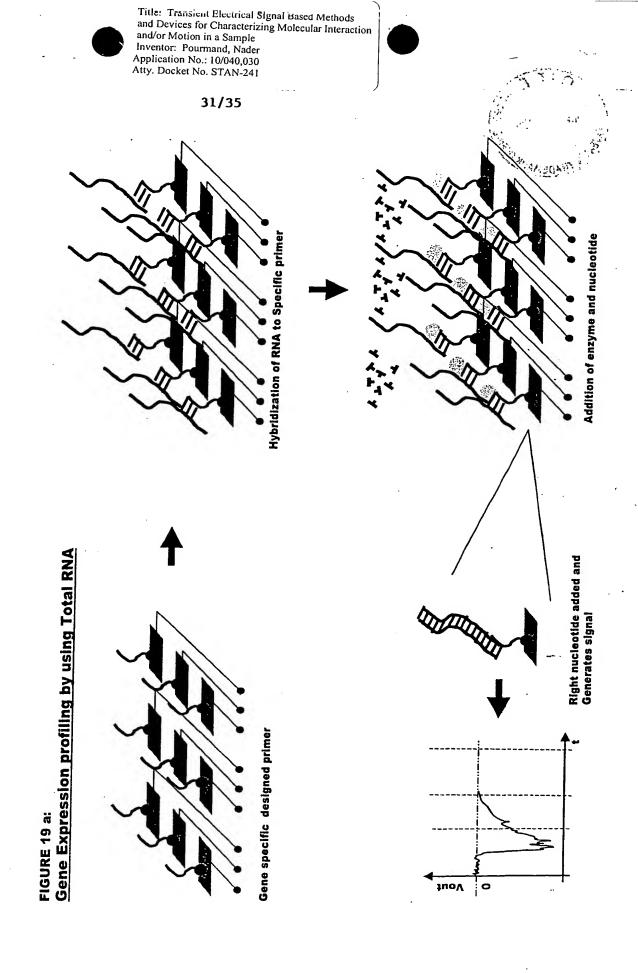
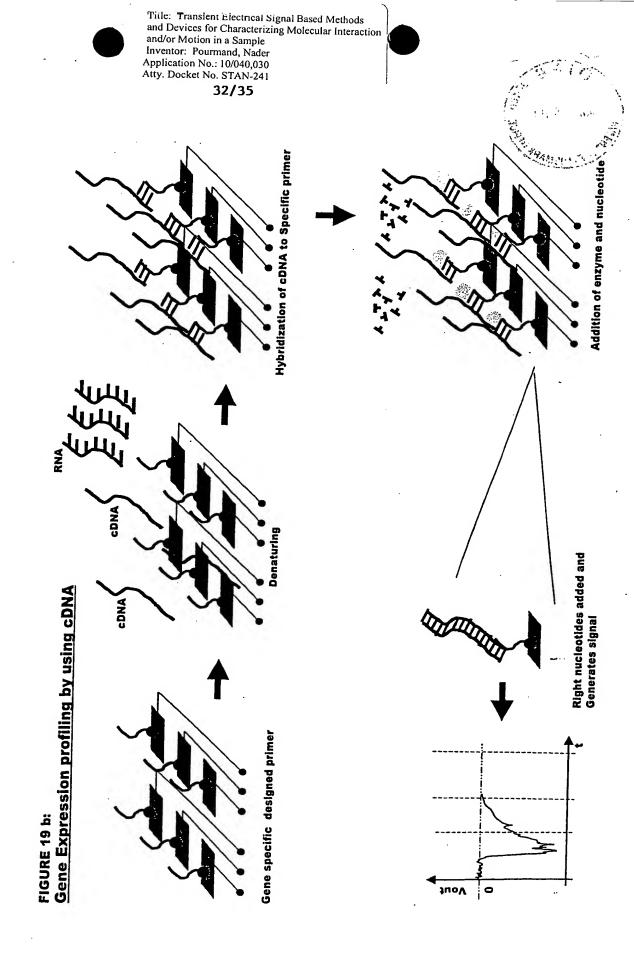


Figure 18D.















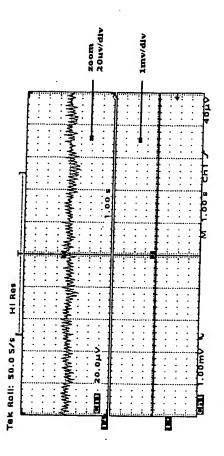


Figure 20





Title: Translent Electrical Signal Based Methods and Devices for Characterizing Molecular Interaction and/or Motion in a Sample

and/or Motion in a Sample Inventor: Pourmand, Nader Application No.: 10/040,030 Atty. Docket No. STAN-241





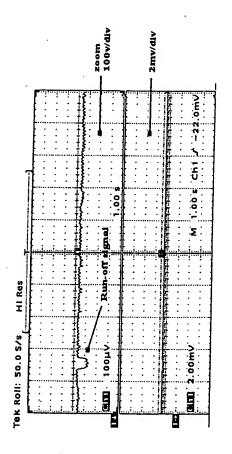


Figure 21

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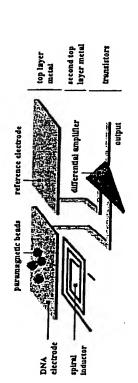


Figure 22